

**INFORMATION DISCLOSURE
STATEMENT BY APPLICANT**

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|---|---|----|---|------------------------|------------------------------|--|--|
| <p style="text-align: right;">Complete if Known</p> | | | | | | | |
| | | | | Application Number | 10758410 10758415 | | |
| | | | | Filing Date | January 16, 2004 | | |
| | | | | First Named Inventor | William S. Brusilow | | |
| | | | | Group Art Unit | 1614 | | |
| Examiner Name | | | | Zohreh Vakili | | | |
| Sheet | 1 | of | 1 | Attorney Docket Number | 2930-109 | | |

NON PATENT LITERATURE DOCUMENTS

| Examiner Initials* | Cite No. ¹ | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published | T ² |
|--------------------|-----------------------|--|----------------|
| /ZV/ | | Hertz et al., "Astrocytes: Glutamate Producers for Neurons", JOURNAL OF NEUROSCIENCE RESEARCH, 57:417-428 (1999). | |
| /ZV/ | | Apostol et al., "A cell-based assay for aggregation inhibitors as therapeutics of polyglutamine-repeat disease and validation in <i>Drosophila</i> "; PROCEEDINGS OF THE NATIONAL ACADEMY OF SCIENCES" 2003 May 13, 100 (10):5950-5955. | |
| /ZV/ | | Swanson et al., "Methionine Sulfoximine Reduces Cortical Infarct Size in Rats After Middle Cerebral Artery Occlusion", STROKE, vol. 21, No. 2, February 1990, pp 322-327. | |
| /ZV/ | | Lamar et al., "The duration of the inhibition of glutamine synthetase by methionine sulfoximine", BIOCHEMICAL PHARMACOLOGY, vol. 17, pp 636-640, 1968 | |
| /ZV/ | | Takahashi et al., "Inhibition of brain glutamine accumulation prevents cerebral edema in hyperammonemic rats", THE AMERICAN PHYSIOLOGICAL SOCIETY, 1991, pgs. H825-H829. | |
| /ZV/ | | Hack et al., "Glufosinate Ammonium-Some aspects of its mode of action in mammals", FD CHEM. TOXIC., vol. 32, no. 5, pp 461-470, 1994. | |
| /ZV/ | | Gill et al., "The crystal structure of phosphinothricin in the active site of glutamine synthetase illuminates the Mechanism of Enzymatic Inhibition", BIOCHEMISTRY 2001, 40, 1903-1912. | |
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| Examiner Signature | /Zohreh Vakili/ (08/24/2007) | Date Considered | /Zohreh Vakili/ (08/24/2007) |
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*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.


¹Unique citation designation number. ²Applicant is to place a check mark here if English language Translation is attached.

* Did not receive


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| INFORMATION DISCLOSURE STATEMENT BY APPLICANT | | | | <i>Complete if Known</i> | |
| | | | | Application Number | New Application 10758415 |
| | | | | Filing Date | Herewith |
| | | | | First Named Inventor | William S. Brusilow |
| | | | | Group Art Unit | 1614 |
| | | | | Examiner Name | Zohreh Vakili |
| Sheet | 1 | of | 1 | Attorney Docket Number | 2930-109 |
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| INFORMATION DISCLOSURE STATEMENT OF APPLICANT  | | | <i>Complete if Known</i> | | |
| | | | Application Number | 10/758,415 | |
| | | | Filing Date | January 16, 2004 | |
| | | | First Named Inventor | William S. Brusilow | |
| | | | Group Art Unit | 1614 | |
| | | | Examiner Name | Zohreh Vakili | |
| Sheet | 1 | of | 2 | Attorney Docket Number | 2930-109 |
| OTHER PRIOR ART - NON PATENT LITERATURE DOCUMENTS | | | | | |
| Examiner Initials* | Cite No. ¹ | Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published | | | T ² |
| /ZV/ | AA | Blei, Andres T., et al., "Ammonia-Induced Brain Edema and Intracranial Hypertension in Rats After Portacaval Anastomosis," <i>Hepatology</i> 19(6): 1437-1444, June 1994. | | | |
| | AB | Brusilow, Saul W. "Inborn Errors of Urea Synthesis," In: Scriver CR, Lloyd Jk, eds. <i>Genetic and Metabolic Disease in Pediatrics</i> . 5: 140-165, London: Butterworths, 1985. | | | |
| | AC | Brusilow, Saul W., et al., "Urea Cycle Disorders: Diagnosis, Pathophysiology, and Therapy," <i>Advances in Pediatrics</i> 43:127-170, 1996. | | | |
| | AD | Butterworth, R.F., "Effects of Hyperammonaemia on Brain Function," <i>J. Inher. Metab. Dis.</i> 21(1):6-20, 1998. | | | |
| | AE | Cordoba, Juan, et al., "Brain Edema and Hepatic Encephalopathy," <i>Seminars in Liver Disease</i> 16(3): 271-280, 1996. | | | |
| | AF | Folbergrova, J., "Free Glutamine Level in the Rat Brain In Vivo After Methionine Sulphoximine Administration," <i>Physiologia Bohemoslovenica</i> 13:21-26, 1963. | | | |
| | AG | Gershoff, S.N., et al., "The Relative Effect of Methionine Sulfoximine on Different Animal Species," <i>J. Nutr.</i> 45:451-458, 1951. | | | |
| | AH | Häussinger, Dieter, et al., "Pathogenesis of Hepatic Encephalopathy," <i>Journal of Gastroenterology and Hepatology</i> 17:S256-S259, 2002. | | | |
| | AI | Hawkins, Richard, et al., "Effect of Reducing Brain Glutamine Synthesis on Metabolic Symptoms of Hepatic Encephalopathy," <i>Journal of Neurochemistry</i> 60(3):1000-1006, 1993. | | | |
| | AJ | Hawkins, Richard, et al., "Hyperammonaemia Does Not Impair Brain Function in the Absence of Net Glutamine Synthesis," <i>Biochem. J.</i> 277:697-703, 1991. | | | |
| | AK | Hirata, Takahiko, et al., "Impaired Pial Arteriolar Reactivity to Hypercapnia During Hyperammonemia Depends on Glutamine Synthesis," <i>Stroke</i> 27(4): 729-736, 1996. | | | |
| | AL | Jonung, Torbjorn, et al., "Methionine Sulfoximine Prevents the Accumulation of Large Neutral Amino Acids in Brain of Hyperammonemic Rats," <i>J. Surgical Research</i> 36:349-353, 1984. | | | |
| | AM | Krakoff, Irwin H., et al., "Effect of Methionine Sulfoximine in Man," <i>J. Pharm. Experimental Ther.</i> 2:599-604, 1961. | | | |
| ↓ | AN | Lamar, C., et al., "The Duration of the Inhibition of Glutamine Synthetase by Methionine Sulfoximine," <i>Biochemical Pharmacology</i> 17:636-642, 1968. | | | |
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| /ZV/ | AO | Master, Sonali, et al., "Cerebral Blood Flow and the Development of Ammonia-Induced Brain Edema in Rats After Portacaval Anastomosis," <i>Hepatology</i> 30(4): 876-880, 1999. | | | |
| | AP | Norenberg, Michael D., et al., "Fine Structural Localization of Glutamine Synthetase in Astrocytes of Rat Brain," <i>Brain Research</i> 161:303-310, 1979. | | | |
| | AQ | Richman, Paul G., et al., "Inhibition of γ-Glutamylcystein Synthetase by L-Methionine-S-Sulfoximine," <i>J. Biological Chemistry</i> 248(19): 6684-6690, 1973. | | | |
| | AR | Rowe, W. Bruce, et al., "Identification of L-Methionine-S-Sulfoximine as the Convulsant Isomer of Methionine Sulfoximine," <i>Proceedings of the National Academy of Sciences</i> 66(2): 500-506, June 1970. | | | |
| | AS | Sellinger, Otto Z., et al., "Methionine Sulfoximine Seizures. VII. the Dissociation of the Convulsant and Glutamine Synthetase Inhibitory Effects," <i>J. Pharmacology & Experimental Therapeutics</i> 161(1): 212-222, 1968. | | | |
| | AT | Sugimoto, Hideyoshi, et al., "Methionine Sulfoximine, A Glutamine Synthetase Inhibitor, Attenuates Increased Extracellular Potassium Activity During Acute Hyperammonemia," <i>Journal of Cerebral Blood Flow & Metabolism</i> , 17:44-49, 1997. | | | |
| | AU | Takahashi, Hideo, et al., "Inhibition of Brain Glutamine Accumulation Prevents Cerebral Edema in Hyperammonemic Rats," <i>American Physiological Society</i> 261:H825-H829, 1991. | | | |
| | AV | Voorhies, Theresa M., "Acute Hyperammonemia in the Young Primate: Physiologic and Neuropathologic Correlates," <i>Pediatric Research</i> 17(12):970-975, 1983. | | | |
| | AW | Wada, John A., et al., "The Susceptibility of Auditory Stimuli of Animals Treated with Methionine Sulfoximine," <i>Experimental Neurology</i> 15:157-165, 1966. | | | |
| | AX | Warren, Kenneth S., et al., "Effect of an Inhibitor of Glutamine Synthesis (Methionine Sulfoximine) on Ammonia Toxicity and Metabolism," <i>J. Lab. & Clin. Med.</i> 64(3): 442-449, 1964. | | | |
| | AY | Watson, Alan J., et al. "Transient Idiopathic Hyperammonaemia in Adults," <i>The Lancet</i> 1271-1274, December 7, 1985. | | | |
| | AZ | Willard-Mack, C.L., et al., "Inhibition of Glutamine Synthetase Reduces Ammonia-Induced Astrocyte Swelling in Rat," <i>Neuroscience</i> 71(2): 589-599, 1996. | | | |
| | BA | Zwingmann, Claudia, et al., "Multinuclear NMR Spectroscopy Studies on NH ₄ Cl-Induced Metabolic Alterations and Detoxification Processes in Primary Astrocytes and Glioma Cells," <i>Dev. Neurosci</i> 20:417-426, 1998. | | | |
| | BB | Apostolakis et al., <i>Brain Research Bulletin</i> , Vol. 23, pp. 257-262 (1989). | | | |
| | BC | Ginefri-Gayet et al., <i>Pharmacology Biochemistry and Behavior</i> , Vol. 43, pp. 173-179 (1992). | | | |
| | BD | Takahashi et al., <i>Circulation Research</i> , 71(5), 1220-1230 (November, 1992). | | | |
| ✓ | BE | Harth et al., <i>J. Exp. Med.</i> , 189(9), 1425-1435 (May 3, 1999). | | | |
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IF

THE UNITED STATES PATENT AND TRADEMARK OFFICE

Appl. No. : 10/758,415
Applicant : William S. Brusilow
Filed : January 16, 2004
TC/A.U. : 1614
Examiner :

Docket No. : 2930-109
Customer No. : 06449
Confirmation No. : 5654

SUPPLEMENTAL INFORMATION DISCLOSURE STATEMENT

Director of the United States Patent
and Trademark Office
P.O. Box 1450
Alexandria, Virginia 22313-1450

Dear Sir:

During a review of the references filed on December 20, 2004, it was noted that the attached reference entitled "Urea Cycle Disorders: Diagnosis, Pathophysiology, and Therapy" was missing several pages. Attached is a complete copy of that reference for the Examiner's review.

Respectfully submitted,

By 
Monica Chin Kitts
Attorney for Applicants
Registration No. 36,105
ROTHWELL, FIGG, ERNST & MANBECK, p.c.
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Washington, D.C. 20005
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RBM/cb